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Complete if Known Substriute for form 1449/PTO **Application Number** 10/735,577 FIRST SUPPLEMENTAL Filing Date 12/12/2003 INFORMATION DISCLOSURE First Named Inventor David Waisman STATEMENT BY APPLICATION Art Unit 1632 (Use as many sheets as necessary) **Examiner Name** Tracy A. Vivlemore Sheet of 1 Attorney Docket Number 101982

| U.S PATENT DOCUMENTS  |              |   |                                |  |   |
|-----------------------|--------------|---|--------------------------------|--|---|
| Examiner<br>Initials* | Cite<br>No.1 | Document Number Number-Kind Code <sup>2 (# Unown)</sup> | Publication Date<br>MM-DD-YYYY | Name of Patentee or Applicant of<br>Cited Document | Pages, Columns, Lines,<br>Where Relevant Passages<br>or Relevant Figures Appeal |
| TV                    | U            | 4,462,980   | 07/31/1984                     | Diedrichsen et al.                                 |   |

| FOREIGN PATENT DOCUMENTS |  |   |                                   |  |  |    |
|--------------------------|--|---|-----------------------------------|--|--|----|
| Examiner<br>Initials*    |  | Foreign Patent Document Country Code <sup>3</sup> -Number <sup>4</sup> -Kind Code <sup>5(*1004**)</sup> | Publication<br>Date<br>MM-DD-YYYY | Name of Patentee or<br>Applicant of Cited Document | Pages, columns, lines,<br>Where Relevant<br>Passages Or Relevant<br>Figures Appear | T⁰ |

| Examiner  | /Tracy Vivlemore/ | Date       |              |
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|                          |    | NON PATENT LITERATURE DOCUMENTS  |  |  |
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|                          |    | BALLAGAMBA et al, "Tyrosine phosphorylation of annexin II tetramer is stimulated by membrane binding." J Biol Chem., 1997, Vol. 272, No. 6, pp. 3195-9.  |  |  |
|                          |    | BROOKS et al., "Ca2+-dependent and phospholipid-independent binding of annexin 2 and annexin 5." Biochem J., 2002, Vol. 367, Pt 3, pp. 895-900.  |  |  |
|                          | X  | CAO et al., "Kringle domains of human angiostatin. Characterization of the anti-proliferative activity on endothelial cells." J Biol Chem., 1996, Vol. 271, No. 46, pp. 29461-7.   |  |  |
| • 1                      | Y  | FALCONE et al., "Macrophage formation of angiostatin during inflammation. A byproduct of the activation of plasminogen." J Biol Chem., 1998, Vol. 273, No. 47, pp 31480-5.   |  |  |
|                          | Z  | GATELY et al., "The mechanism of cancer-mediated conversion of plasminogen to the angiogenesis inhibitor angiostatin." Proc Natl Acad Sci U S A, 1997, Vol. 94, No. 20, pp 10868-72.   |  |  |
|                          | AA | JOHNSSON et al., "Alkylation of cysteine 82 of p11 abolishes the complex formation with the tyrosine-protein kinase substrate p36 (annexin 2, calpactin 1, lipocortin 2)." J Biol Chem., 1990, Vol. 265, No. 24, pp. 14464-8.                                      |  |  |
|                          | AB | KASSAM et al. "Characterization of the heparin binding properties of annexin II tetramer." J Biol Chem., 1997, Vol. 272, No. 24, pp. 15093-100.  |  |  |
|                          | AC | KWON et al., "p22 is a novel plasminogen fragment with antiangiogenic activity." Biochemistry, 2001, Vol. 40, No. 44, pp. 13246-53.  |  |  |
|                          | AD | Soff, "Angiostatin and angiostatin-related proteins." Cancer Metastasis Rev., 2000, Vol. 19, No. (1-2), pp. 97-107.  |  |  |
|                          | AE | STATHAKIS et al., "Angiostatin formation involves disulfide bond reduction and proteolysis in kringle 5 of plasmin." J Biol Chem., 1999, Vol. 274, No. 13, pp. 8910-6.   |  |  |
| V                        | AF | TERATANI et al., "Induced transcriptional expression of calcium-binding protein S100A1 and S100A10 genes in human renal cell carcinoma." Cancer Lett., 2002, Vol. 175, No. 1, pp. 71-7.  |  |  |

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